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Substitute for form 1449 PTO

## INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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Sheet

1

of

6

**Complete if Known**

<b>Application Number</b>	<b>10/783,986</b>
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<b>Filing Date</b>	<b>February 19, 2004</b>
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<b>First Named Inventor</b>	<b>Ronit Satchi-Fainaro et al.</b>
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Art Unit	1614
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<b>Examiner Name</b>	<b>To be assigned</b>
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Attorney Docket Number	701039-52584-CIP
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## U.S. PATENT DOCUMENTS

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**FOREIGN PATENT DOCUMENTS**

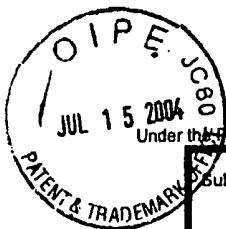
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				Art Unit	1614
				Examiner Name	To be assigned
Sheet	2	of	6	Attorney Docket Number	701039-52584-CIP

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
VB	C1	Folkman, J., Angiogenesis. in <i>Harrison's Textbook of Internal Medicine</i> (eds. Braunwald, E. et al.) 517-530 (McGraw Hill, New York, 2001).	
	C2	Hanahan, D. et al., Patterns and emerging mechanisms of the angiogenic switch during tumorigenesis, <i>Cell</i> , 86:353-64 (1996).	
	C3	Volpert, O.V. et al., Id1 regulates angiogenesis through transcriptional repression of thrombospondin-1, <i>Cancer Cell</i> , 2:473-483 (2002).	
	C4	Folkman, J., Tumor angiogenesis, <i>Cancer Medicine</i> (eds. Holland, J. et al.), pp. 132-152 (B. C. Decker Inc., Ontario, Canada, 2000).	
	C5	Lyden, D. et al., Id1 and Id3 are required for neurogenesis, angiogenesis and vascularization of tumour xenografts, <i>Nature</i> , 401:670-677 (1999).	
	C6	Streit, M. et al., Thrombospondin-2: a potent endogenous inhibitor of tumor growth and angiogenesis, <i>Proc Natl. Acad. Sci. USA</i> , 96:14888-14893 (1999).	
	C7	Chin, L. et al., Essential role for oncogenic Ras in tumour maintenance, <i>Nature</i> , 400:468-472 (1999).	
	C8	Tabone, M.D. et al., Are basic fibroblast growth factor and vascular endothelial growth factor prognostic indicators in pediatric patients with malignant solid tumors?, <i>Clinical Cancer Res.</i> , 7:538-543 (2001).	
	C9	Yao, Y. et al., Prognostic value of vascular endothelial growth factor and its receptors Flt-1 and Flk-1 in astrocytic tumours, <i>Acta Neurochir (Wien)</i> , 143:159-66 (2001).	
	VM	C10	Yuan, A. et al., Aberrant p53 expression correlates with expression of vascular endothelial growth factor mRNA and interleukin-8 mRNA and neoangiogenesis in non-small-cell lung cancer, <i>J. Clinical Oncology</i> , 20:900-910 (2002).

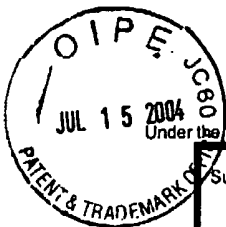
Examiner Signature	V. Balasubramaniam	Date Considered	9/30/04
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			First Named Inventor	Ronit Satchi-Fainaro et al.	
			Art Unit	1614	
			Examiner Name	To be assigned	
Sheet	3	of	6	Attorney Docket Number	701039-52584-CIP

✓	C11	Ingber, D. et al., Synthetic analogues of fumagillin that inhibit angiogenesis and suppress tumour growth, <i>Nature</i> , 348:555-557 (1990).	
	C12	Antoine, N. et al., AGM-1470, a potent angiogenesis inhibitor, prevents the entry of normal but not transformed endothelial cells into the G <sub>1</sub> phase of the cell cycle, <i>Cancer Res.</i> , 54:2073-2076 (1994).	
	C13	Kudelka, A.P. et al., Complete remission of metastatic cervical cancer with the angiogenesis inhibitor TNP-470, <i>N. Engl. J. Med.</i> , 338:991-2 (1998).	
	C14	Kudelka, A.P. et al., A phase I study of TNP-470 administered to patients with advanced squamous cell cancer of the cervix, <i>Clinical Cancer Res.</i> , 3:1501-1505 (1997).	
	C15	Bhargava, P. et al., A Phase I and pharmacokinetic study of TNP-470 administered weekly to patients with advanced cancer, <i>Clinical Cancer Res.</i> , 5:1989-1995 (1999).	
	C16	Herbst, R.S. et al., Safety and pharmacokinetic effects of TNP-470, an angiogenesis inhibitor, combined with paclitaxel in patients with solid tumors: evidence for activity in non-small-cell lung cancer, <i>J. Clinical Oncol.</i> , 20:4440-4447 (2002).	
	C17	Kim, E.S. et al., Angiogenesis inhibitors in lung cancer. <i>Curr. Oncol. Rep.</i> , 4:325-333 (2002).	
	C18	Stadler, W.M. et al., Multi-institutional study of the angiogenesis inhibitor TNP-470 in metastatic renal carcinoma, <i>J. Clinical Oncol.</i> , 17:2541-2545 (1999).	
	C19	Logothetis, C.J. et al., Phase I trial of the angiogenesis inhibitor TNP-470 for progressive androgen-independent prostate cancer. <i>Clinical Cancer Res.</i> , 7:1198-1203 (2001).	
	C20	Rupnick, M.A. et al., Adipose tissue mass can be regulated through the vasculature, <i>Proc. Natl. Acad. Sci. U.S.A.</i> , 99:10730-10735 (2002).	
✓	C21	Schoof, D.D. et al., The influence of angiogenesis inhibitor AGM-1470 on immune system status and tumor growth in vitro, <i>Int. J. Cancer</i> , 55:630-635 (1993).	

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Sheet	4	of	6	Attorney Docket Number	701039-52584-CIP

M	C22	Nagabuchi, E. et al., TNP-470 antiangiogenic therapy for advanced murine neuroblastoma, <i>J. Pediatric Surg.</i> , 32:287-93 (1997).	
	C23	Rihova, B. et al., Biocompatibility of N-(2-hydroxypropyl) methacrylamide copolymers containing adriamycin. Immunogenicity, and effect on haematopoietic stem cells in bone marrow in vivo and mouse splenocytes and human peripheral blood lymphocytes in vitro, <i>Biomaterials</i> , 10:335-342. (1989).	
	C24	Seymour, L.W. et al., The pharmacokinetics of polymer-bound adriamycin, <i>Biochem. Pharmacol.</i> , 39:1125-1131 (1990).	
	C25	Maeda, H. et al., Tumor vascular permeability and the EPR effect in macromolecular therapeutics: a review, <i>J. Controlled Release</i> , 65:271-284 (2000).	
	C26	Duncan, R. et al., Preclinical toxicology of a novel polymeric antitumour agent: HPMA copolymer-doxorubicin (PK1), <i>Human and Exp. Toxicology</i> , 17:93-104 (1998).	
	C27	Satchi-Fainaro, R., Targeting tumor vasculature: Reality or a dream?. <i>J. Drug Targeting</i> , 10:529-533 (2002).	
	C28	Duncan, R. et al., Polymers containing enzymatically degradable bonds, 7. Design of oligopeptide side chains in poly [N-(2-hydroxypropyl)methacrylamide] copolymers to promote efficient degradation by lysosomal enzymes, <i>Makromol. Chem.</i> , 184:1997-2008 (1983).	
	C29	Foekens, J.A. et al., Prognostic significance of cathepsins B and L in primary human breast cancer. <i>J. Clinical Oncol.</i> , 16:1013-1021 (1998).	
	C30	Gianasi, E. et al.. HPMA copolymer platinates as novel antitumour agents: in vitro properties, pharmacokinetics and antitumour activity in vivo, <i>Eur. J. Cancer</i> , 35:994-1002 (1999).	
	C31	Kusaka, M. et al. Cytostatic inhibition of endothelial cell growth by the angiogenesis inhibitor TNP-470 (AGM-1470), <i>Br. J. Cancer</i> . 69:212-216 (1994).	
M	C32	Greene, A.K. et al., Endothelial-directed hepatic regeneration after partial hepatectomy, <i>Ann. Surg.</i> , 237:530-535 (2003)	

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Sheet	5	of	6	Attorney Docket Number	701039-52584-CIP

M	C33	Drixler, T.A. et al., Liver regeneration is an angiogenesis- associated phenomenon, <i>Ann. Surg.</i> , 236:703-712 (2002).	
	C34	Klein, S.A. et al., Angiogenesis inhibitor TNP-470 inhibits murine cutaneous wound healing, <i>J. Surg. Res.</i> , 82:268-274 (1999).	
	C35	Whalen, C.T. et al., Assay of TNP-470 and its two major metabolites in human plasma by high-performance liquid chromatography-mass spectrometry, <i>J. Chromatographic Sci.</i> , 40:214-218 (2002).	
	C36	Brocchini, S. et al., Polymer-Drug conjugates: drug release from pendent linkers. in <i>Encyclopaedia of controlled release</i> (ed. Mathiovitz, E.) 786-816 (New York: Wiley, 1999).	
	C37	Duncan, R. et al., Polymer-drug conjugates, PDEPT and PELT: basic principles for design and transfer from the laboratory to clinic, <i>J. Controlled Release</i> , 74:135-146 (2001).	
	C38	Vasey, P.A. et al., Phase I clinical and pharmacokinetic study of PK1 [N-(2-hydroxypropyl)methacrylamide copolymer doxorubicin]: first member of a new class of chemotherapeutic agents-drug-polymer conjugates, Cancer Research Campaign Phase I/II Committee, <i>Clinical Cancer Res.</i> , 5:83-94 (1999).	
	C39	Seymour, L.W. et al., Tumour tropism and anti-cancer efficacy of polymer-based doxorubicin prodrugs in the treatment of subcutaneous murine B16F10 melanoma, <i>Br. J. Cancer</i> , 70:636-641 (1994).	
	C40	Dvorak, H.F. et al., Identification and characterization of the blood vessels of solid tumors that are leaky to circulating macromolecules. <i>Am. J. Pathology</i> , 133:95-109 (1988).	
	C41	Griffith, E.C. et al., Methionine aminopeptidase (type 2) is the common target for angiogenesis inhibitors AGM-1470 and ovalicin, <i>Chem. and Biol.</i> , 4, 461-471 (1997).	
	C42	Auerbach, R. et al., Angiogenesis assays: problems and pitfalls, <i>Cancer Metastasis Rev.</i> , 19:167-172 (2000).	
M	C43	Seymour, L.W. et al., Hepatic drug targeting: phase I evaluation of polymer-bound doxorubicin., <i>J. Clinical Oncol.</i> , 20:1668-1676 (2002).	

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W	C44	Francis, G.E. et al., PEG-modified proteins. in <i>Stability of Proteins Pharmaceuticals (Part B)</i> (ed. Ahem TJ, M.M.) 235-263 (Plenum Press, New York, 1992).	
	C45	Ho, D.H. et al., Clinical pharmacology of polyethylene glycol-L-asparaginase, <i>Drug Metabolism Disposition</i> , 14:349-352 (1986).	
	C46	O'Reilly, M.S. et al., Angiostatin: a novel angiogenesis inhibitor that mediates the suppression of metastases by a Lewis lung carcinoma, <i>Cell</i> , 79:315-328 (1994).	
	C47	Folkman, J. et al., Long-term culture of capillary endothelial cells, <i>Proc. Natl. Acad. Sci. USA</i> , 76:5217-5221 (1979).	
	C48	Waynforth, H.B. Routes and methods of administration, Intracerebral injection. in <i>Experimental and Surgical technique in the rat</i> , Vol. 2.9 34-36 (Academic Press, London, 1980).	
	C49	Bhargava, P. et al., A Phase I and pharmacokinetic study of TNP-470 administered weekly to patients with advanced cancer, <i>Clinical Cancer Res.</i> , 5:1989-1995 (1999).	
	C50	Seymour, L.W. et al., The pharmacokinetics of polymer-bound adriamycin, <i>Biochemical Pharmacology</i> , 39:1125-1131 (1990).	
	C51	Yeh, J.R. et al., The antiangiogenic agent TNP-470 requires p53 and p21 <sup>CIP/WAF</sup> for endothelial cell growth arrest, <i>Proc. Natl. Acad. Sci. USA</i> , 97:12782-12787 (2000).	
	C52	Zhang, Y. et al., Cell cycle inhibition by the anti-angiogenic agent TNP-470 is mediated by p53 and p21 <sup>WAF1/CIP1</sup> , <i>Proc. Natl. Acad. Sci. USA</i> , 97:6427-6432 (2000).	
	C53	Seymour, L.W. et al., N-(2-hydroxypropyl) methacrylamide copolymers targeted to the hepatocyte galactose-receptor: pharmacokinetics in DBA <sub>2</sub> mice, <i>Br. J. Cancer</i> , 63:859-866 (1991).	
M	C54	Folkman, J. Tumor angiogenesis. in <i>Accomplishments in cancer research</i> (eds. Wells, S.J. & Sharp, P.) 32-44 (Lippincott Williams & Wilkins, New York, 1998)	

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